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CD-Rom

PATENT
ATTORNEY'S DOCKET NO.: NBG-109

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants: Aldo Salimbeni et al.
Serial No.: 10/537,731
Filed: June 6, 2005
For: PROCESS FOR THE PREPARATION OF BICYCLIC
PEPTIDE COMPOUNDS
Examiner: Not Assigned
Art Unit: Not Assigned

CERTIFICATE OF MAILING

I hereby certify that the following correspondence is being deposited with the United States Postal service as first class mail in an envelope addressed to the Commissioner for Patents, PO Box 1450, Alexandria, VA 22313-1450 on February 22, 2007.


Mark D. Lorusso

Commissioner for Patents
PO BOX 1450
Alexandria, VA 22313-1450

RESPONSE TO STIC BIOTECHNOLOGY SYSTEMS BRANCH
RAW SEQUENCE LISTING ERROR REPORT

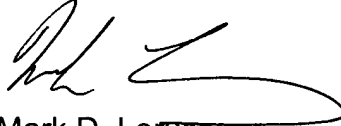
Sir:

This responds to the Raw Sequence Listing Error Report processed July 3, 2006. On February 21, 2007, applicants' counsel of record spoke with Anne-Marie Corrigan in the STIC Biotechnology Systems Branch with respect to the Raw Sequence Report and was faxed a copy of the report not previously received. Ms. Corrigan explained the substantive errors that could not be identified with the CHECKER software.

Enclosed herewith are two copies of the corrected Sequence Listing in computer readable format. Another copy of the corrected Sequence Listing in paper form is also enclosed. The content of the Sequence Listing information recorded in computer

readable form is identical to the written sequence listing and does not include any new matter. A courtesy copy of the Raw Sequence Listing Error Report is also enclosed to facilitate processing of this response.

Respectfully submitted,

A handwritten signature in black ink, appearing to read 'Mark D. Lorusso', with a long horizontal flourish extending to the right.

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Reg. No. 41,955
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Portsmouth, NH 03801
Tel: (603) 427-0070
Attorneys for Applicants

Docket No.: NBG-109

Date: February 22, 2007

**STIC Biotechnology Systems Branch****RAW SEQUENCE LISTING**
ERROR REPORT

The Biotechnology Systems Branch of the Scientific and Technical Information Center (STIC) detected errors when processing the following computer readable form:

Application Serial Number: 10/537,731
Source: PTA
Date Processed by STIC: 7/3/06

THE ATTACHED PRINTOUT EXPLAINS DETECTED ERRORS.

PLEASE FORWARD THIS INFORMATION TO THE APPLICANT BY EITHER:

- 1) INCLUDING A COPY OF THIS PRINTOUT IN YOUR NEXT COMMUNICATION TO THE APPLICANT, WITH A NOTICE TO COMPLY or,
- 2) TELEPHONING APPLICANT AND FAXING A COPY OF THIS PRINTOUT, WITH A NOTICE TO COMPLY

FOR CRF SUBMISSION AND PATENTIN SOFTWARE QUESTIONS, PLEASE CONTACT MARK SPENCER, TELEPHONE: 571-272-2510; FAX: 571-273-0221

TO REDUCE ERRORED SEQUENCE LISTINGS, PLEASE USE THE **CHECKER** **VERSION 4.4.0 PROGRAM**, ACCESSIBLE THROUGH THE U.S. PATENT AND TRADEMARK OFFICE WEBSITE. SEE BELOW FOR ADDRESS:

<http://www.uspto.gov/web/offices/pac/checker/chkrnote.htm>

Applicants submitting genetic sequence information electronically on diskette or CD-Rom should be aware that there is a possibility that the disk/CD-Rom may have been affected by treatment given to all incoming mail.

Please consider using alternate methods of submission for the disk/CD-Rom or replacement disk/CD-Rom.

Any reply including a sequence listing in electronic form should NOT be sent to the 20231 zip code address for the United States Patent and Trademark Office, and instead should be sent via the following to the indicated addresses:

1. EFS-Bio (<<http://www.uspto.gov/ebc/efs/downloads/documents.htm>> , EFS Submission User Manual - ePAVE)
2. U.S. Postal Service: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450
3. Hand Carry, Federal Express, United Parcel Service, or other delivery service (EFFECTIVE 01/14/05):
U.S. Patent and Trademark Office, Mail Stop Sequence, Customer Window, Randolph Building, 401 Dulany Street, Alexandria, VA 22314

Revised 01/10/06

Raw Sequence Listing Error Summary

ERROR DETECTED	SUGGESTED CORRECTION	SERIAL NUMBER: 10/537,731
ATTN: NEW RULES CASES: PLEASE DISREGARD ENGLISH "ALPHA" HEADERS, WHICH WERE INSERTED BY PTO SOFTWARE		
1 ____ Wrapped Nucleics Wrapped Aminos	The number/text at the end of each line "wrapped" down to the next line. This may occur if your file was retrieved in a word processor after creating it. Please adjust your right margin to .3; this will prevent "wrapping."	
2 ____ Invalid Line Length	The rules require that a line not exceed 72 characters in length. This includes white spaces.	
3 ____ Misaligned Amino Numbering	The numbering under each 5 th amino acid is misaligned. Do not use tab codes between numbers; use space characters, instead.	
4 ____ Non-ASCII	The submitted file was not saved in ASCII(DOS) text, as required by the Sequence Rules. Please ensure your subsequent submission is saved in ASCII text.	
5 ____ Variable Length	Sequence(s) ____ contain n's or Xaa's representing more than one residue. Per Sequence Rules, each n or Xaa can only represent a single residue. Please present the maximum number of each residue having variable length and indicate in the <220>-<223> section that some may be missing.	
6 ____ PatentIn 2.0 "bug"	A "bug" in PatentIn version 2.0 has caused the <220>-<223> section to be missing from amino acid sequences(s) _____. Normally, PatentIn would automatically generate this section from the previously coded nucleic acid sequence. Please manually copy the relevant <220>-<223> section to the subsequent amino acid sequence. This applies to the mandatory <220>-<223> sections for Artificial or Unknown sequences.	
7 ____ Skipped Sequences (OLD RULES)	Sequence(s) ____ missing. If intentional, please insert the following lines for each skipped sequence: (2) INFORMATION FOR SEQ ID NO:X: (insert SEQ ID NO where "X" is shown) (i) SEQUENCE CHARACTERISTICS: (Do not insert any subheadings under this heading) (xi) SEQUENCE DESCRIPTION:SEQ ID NO:X: (insert SEQ ID NO where "X" is shown) This sequence is intentionally skipped Please also adjust the "(ii) NUMBER OF SEQUENCES:" response to include the skipped sequences.	
8 ____ Skipped Sequences (NEW RULES)	Sequence(s) ____ missing. If intentional, please insert the following lines for each skipped sequence. <210> sequence id number <400> sequence id number 000	
9 ____ Use of n's or Xaa's (NEW RULES)	Use of n's and/or Xaa's have been detected in the Sequence Listing. Per 1.823 of Sequence Rules, use of <220>-<223> is MANDATORY if n's or Xaa's are present. In <220> to <223> section, please explain location of n or Xaa, and which residue n or Xaa represents.	
10 ____ Invalid <213> Response	Per 1.823 of Sequence Rules, the only valid <213> responses are: Unknown, Artificial Sequence, or scientific name (Genus/species). <220>-<223> section is required when <213> response is Unknown or is Artificial Sequence. (see item 11 below)	
11 ____ Use of <220>	Sequence(s) ____ missing the <220> "Feature" and associated numeric identifiers and responses. Use of <220> to <223> is MANDATORY if <213> "Organism" response is "Artificial Sequence" or "Unknown." Please explain source of genetic material in <220> to <223> section or use "chemically synthesized" as explanation. (See "Federal Register," 06/01/1998, Vol. 63, No. 104, pp. 29631-32), also Sec. 1.823 of Sequence Rules	
12 ____ PatentIn 2.0 "bug"	Please do not use "Copy to Disk" function of PatentIn version 2.0. This causes a corrupted file, resulting in missing mandatory numeric identifiers and responses (as indicated on raw sequence listing). Instead, please use "File Manager" or any other manual means to copy file to floppy disk.	
13 ____ Misuse of n/Xaa	"n" can only represent a single nucleotide; "Xaa" can only represent a single amino acid	



PCT

RAW SEQUENCE LISTING

PATENT APPLICATION: US/10/537,731

DATE: 07/03/2006

TIME: 11:56:39

Input Set : E:\3874 PTUS sequence listing.txt
Output Set: N:\CRF4\07032006\J537731.raw

3 <110> APPLICANT: SALIMBENI, Aldo et al
5 <120> TITLE OF INVENTION: Process for the preparation of bicyclic hexa-peptide
nepadutant

7 <130> FILE REFERENCE: 3874PTUS
C--> 9 <140> CURRENT APPLICATION NUMBER: US/10/537,731
C--> 10 <141> CURRENT FILING DATE: 2005-06-06
12 <150> PRIOR APPLICATION NUMBER: FI2002A000239
13 <151> PRIOR FILING DATE: 2002-06-12
15 <160> NUMBER OF SEQ ID NOS: 11
17 <170> SOFTWARE: PatentIn version 3.3
19 <210> SEQ ID NO: 1
20 <211> LENGTH: 5
21 <212> TYPE: PRT
22 <213> ORGANISM: pentapeptide
25 <220> FEATURE:
26 <221> NAME/KEY: BINDING
27 <222> LOCATION: (1)..(1)
28 <223> OTHER INFORMATION: Asp is bound to a benzyloxycarbonyl group
30 <220> FEATURE:
31 <221> NAME/KEY: MISC_FEATURE
32 <222> LOCATION: (4)..(4)
33 <223> OTHER INFORMATION: X is Dpr (i.e. 2,3-diaminopropionic acid)
35 <220> FEATURE:
36 <221> NAME/KEY: MOD_RES
37 <222> LOCATION: (5)..(5)
38 <223> OTHER INFORMATION: METHYLATION
40 <400> SEQUENCE: 1

W--> 42 Asp Trp Phe Xaa Leu

43 1. 5
46 <210> SEQ ID NO: 2
47 <211> LENGTH: 5
48 <212> TYPE: PRT
49 <213> ORGANISM: Artificial Sequence
51 <220> FEATURE: never has a vsp.
52 <223> OTHER INFORMATION: cyclic pentapeptide
55 <220> FEATURE:
56 <221> NAME/KEY: BINDING
57 <222> LOCATION: (1)..(1)
58 <223> OTHER INFORMATION: Asp is bound to a benzyloxycarbonyl group
60 <220> FEATURE:
61 <221> NAME/KEY: SITE
62 <222> LOCATION: (1)..(4)
63 <223> OTHER INFORMATION: Asp and Dpr are bound together to form a cycle
65 <220> FEATURE:

*see
pg 1, 6*
**Does Not Comply
Corrected Diskette Needed**

*invalid response -
see item 10 on
Error Summary Sheet*

RAW SEQUENCE LISTING

DATE: 07/03/2006

PATENT APPLICATION: US/10/537,731

TIME: 11:56:39

Input Set : E:\3874 PTUS sequence listing.txt

Output Set: N:\CRF4\07032006\J537731.raw

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66 <221> NAME/KEY: MISC_FEATURE
67 <222> LOCATION: (4)..(4)
68 <223> OTHER INFORMATION: X is Dpr (i.e. 2,3-diaminopropionic acid)
70 <220> FEATURE:
71 <221> NAME/KEY: MOD_RES
72 <222> LOCATION: (5)..(5)
73 <223> OTHER INFORMATION: METHYLATION
75 <400> SEQUENCE: 2
W--> 77 Asp Trp Phe Xaa Leu
78 1 5
81 <210> SEQ ID NO: 3
82 <211> LENGTH: 5
83 <212> TYPE: PRT
84 <213> ORGANISM: Artificial Sequence
86 <220> FEATURE:
87 <223> OTHER INFORMATION: cyclic pentapeptide
90 <220> FEATURE:
91 <221> NAME/KEY: BINDING
92 <222> LOCATION: (1)..(1)
93 <223> OTHER INFORMATION: Asp is bound to a benzyloxycarbonyl group
95 <220> FEATURE:
96 <221> NAME/KEY: SITE
97 <222> LOCATION: (1)..(4)
98 <223> OTHER INFORMATION: Asp and Dpr are bound together to form a cycle
100 <220> FEATURE:
101 <221> NAME/KEY: MISC_FEATURE
102 <222> LOCATION: (4)..(4)
103 <223> OTHER INFORMATION: X is Dpr (i.e. 2,3-diaminopropionic acid)
105 <400> SEQUENCE: 3
W--> 107 Asp Trp Phe Xaa Leu
108 1 5
111 <210> SEQ ID NO: 4
112 <211> LENGTH: 6
113 <212> TYPE: PRT
114 <213> ORGANISM: Artificial Sequence
116 <220> FEATURE:
117 <223> OTHER INFORMATION: cyclic hexapeptide
120 <220> FEATURE:
121 <221> NAME/KEY: BINDING
122 <222> LOCATION: (1)..(1)
123 <223> OTHER INFORMATION: Asp is bound to a benzyloxycarbonyl group and to a tert-
butyl
124 group
126 <220> FEATURE:
127 <221> NAME/KEY: SITE
128 <222> LOCATION: (2)..(5)
129 <223> OTHER INFORMATION: Asp and Dpr are bound together to form a cycle
131 <220> FEATURE:
132 <221> NAME/KEY: MISC_FEATURE
133 <222> LOCATION: (5)..(5)

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RAW SEQUENCE LISTING

DATE: 07/03/2006

PATENT APPLICATION: US/10/537,731

TIME: 11:56:39

Input Set : E:\3874 PTUS sequence listing.txt

Output Set: N:\CRF4\07032006\J537731.raw

134 <223> OTHER INFORMATION: X is Dpr (i.e. 2,3-aminopropionic acid)
136 <400> SEQUENCE: 4
W--> 138 Asp Asp Trp Phe Xaa Leu
139 1 5
142 <210> SEQ ID NO: 5
143 <211> LENGTH: 6
144 <212> TYPE: PRT
145 <213> ORGANISM: Artificial Sequence
147 <220> FEATURE:
148 <223> OTHER INFORMATION: bicyclic hexapeptide
151 <220> FEATURE:
152 <221> NAME/KEY: SITE
153 <222> LOCATION: (1)..(6)
154 <223> OTHER INFORMATION: Asp and Leu are bound together to form a cycle
156 <220> FEATURE:
157 <221> NAME/KEY: BINDING
158 <222> LOCATION: (1)..(1)
159 <223> OTHER INFORMATION: Asp is bound to a tert-butyl group
161 <220> FEATURE:
162 <221> NAME/KEY: SITE
163 <222> LOCATION: (2)..(4)
164 <223> OTHER INFORMATION: Asp and Dpr are bound together to form a cycle
166 <220> FEATURE:
167 <221> NAME/KEY: MISC_FEATURE
168 <222> LOCATION: (5)..(5)
169 <223> OTHER INFORMATION: X is Dpr (i.e. 2,3-diaminopropionic acid)
171 <400> SEQUENCE: 5
W--> 173 Asp Asp Trp Phe Xaa Leu
174 1 5
177 <210> SEQ ID NO: 6
178 <211> LENGTH: 6
179 <212> TYPE: PRT
180 <213> ORGANISM: Artificial Sequence
182 <220> FEATURE:
183 <223> OTHER INFORMATION: bicyclic hexapeptide
186 <220> FEATURE:
187 <221> NAME/KEY: SITE
188 <222> LOCATION: (1)..(6)
189 <223> OTHER INFORMATION: Asp and Leu are bound together to form a cycle
191 <220> FEATURE:
192 <221> NAME/KEY: SITE
193 <222> LOCATION: (2)..(5)
194 <223> OTHER INFORMATION: Asp and Dpr are bound together to form a cycle
196 <220> FEATURE:
197 <221> NAME/KEY: MISC_FEATURE
198 <222> LOCATION: (5)..(5)
199 <223> OTHER INFORMATION: X is Dpr (i.e. 2,3-diaminopropionic acid)
201 <400> SEQUENCE: 6
W--> 203 Asp Asp Trp Phe Xaa Leu

RAW SEQUENCE LISTING

DATE: 07/03/2006

PATENT APPLICATION: US/10/537,731

TIME: 11:56:39

Input Set : E:\3874 PTUS sequence listing.txt

Output Set: N:\CRF4\07032006\J537731.raw

```
204 1          5
207 <210> SEQ ID NO: 7
208 <211> LENGTH: 6
209 <212> TYPE: PRT
210 <213> ORGANISM: Artificial Sequence
212 <220> FEATURE:
213 <223> OTHER INFORMATION: bicyclic glycopeptide
216 <220> FEATURE:
217 <221> NAME/KEY: SITE
218 <222> LOCATION: (1)..(6)
219 <223> OTHER INFORMATION: Asp and Leu are bound together to form a cycle
221 <220> FEATURE:
222 <221> NAME/KEY: CARBOHYD
223 <222> LOCATION: (1)..(1)
224 <223> OTHER INFORMATION: Asp is bound to
225     2-acetamide-3,4,6-tri-O-acetyl-2-deoxy-beta-D-glucopyranosylamine
227 <220> FEATURE:
228 <221> NAME/KEY: SITE
229 <222> LOCATION: (2)..(5)
230 <223> OTHER INFORMATION: Asp and Dpr are bound together to form a cycle
232 <220> FEATURE:
233 <221> NAME/KEY: MISC_FEATURE
234 <222> LOCATION: (5)..(5)
235 <223> OTHER INFORMATION: X is Dpr (i.e. 2,3-diaminopropionic acid)
237 <400> SEQUENCE: 7
W--> 239 Asp Asp Trp Phe Xaa Leu
240 1          5
243 <210> SEQ ID NO: 8
244 <211> LENGTH: 6
245 <212> TYPE: PRT
246 <213> ORGANISM: Artificial Sequence
248 <220> FEATURE:
249 <223> OTHER INFORMATION: bicyclic glycopeptide
252 <220> FEATURE:
253 <221> NAME/KEY: SITE
254 <222> LOCATION: (1)..(6)
255 <223> OTHER INFORMATION: Asp and Leu are bound together to form a cycle
257 <220> FEATURE:
258 <221> NAME/KEY: CARBOHYD
259 <222> LOCATION: (1)..(1)
260 <223> OTHER INFORMATION: Asp is bound to 2-acetamide-2-deoxy-beta-D-
glucopyranosylamine
262 <220> FEATURE:
263 <221> NAME/KEY: SITE
264 <222> LOCATION: (2)..(5)
265 <223> OTHER INFORMATION: Asp and Dpr are bound together to form a cycle
267 <220> FEATURE:
268 <221> NAME/KEY: MISC_FEATURE
269 <222> LOCATION: (5)..(5)
270 <223> OTHER INFORMATION: X is Dpr (i.e. 2,3-diaminopropionic acid)
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RAW SEQUENCE LISTING

DATE: 07/03/2006

PATENT APPLICATION: US/10/537,731

TIME: 11:56:39

Input Set : E:\3874 PTUS sequence listing.txt

Output Set: N:\CRF4\07032006\J537731.raw

272 <400> SEQUENCE: 8
W--> 274 Asp Asp Trp Phe Xaa Leu
275 1 5
278 <210> SEQ ID NO: 9
279 <211> LENGTH: 4
280 <212> TYPE: PRT
281 <213> ORGANISM: Artificial Sequence
283 <220> FEATURE:
284 <223> OTHER INFORMATION: tetrapeptide
287 <220> FEATURE:
288 <221> NAME/KEY: BINDING
289 <222> LOCATION: (1)..(1)
290 <223> OTHER INFORMATION: Trp is bound to a benzyloxycarbonyl group
292 <220> FEATURE:
293 <221> NAME/KEY: MISC_FEATURE
294 <222> LOCATION: (3)..(3)
295 <223> OTHER INFORMATION: X is Dpr (i.e. 2,3-diaminopropionic acid)
297 <220> FEATURE:
298 <221> NAME/KEY: BINDING
299 <222> LOCATION: (3)..(3)
300 <223> OTHER INFORMATION: Dpr is bound to a tert-butoxycarbonyl group
302 <220> FEATURE:
303 <221> NAME/KEY: MOD_RES
304 <222> LOCATION: (4)..(4)
305 <223> OTHER INFORMATION: METHYLATION
307 <400> SEQUENCE: 9
W--> 309 Trp Phe Xaa Leu
310 1
313 <210> SEQ ID NO: 10
314 <211> LENGTH: 4
315 <212> TYPE: PRT
316 <213> ORGANISM: Artificial Sequence
318 <220> FEATURE:
319 <223> OTHER INFORMATION: tetrapeptide
322 <220> FEATURE:
323 <221> NAME/KEY: MISC_FEATURE
324 <222> LOCATION: (3)..(3)
325 <223> OTHER INFORMATION: X is Dpr (i.e. 2,3-diaminopropionic acid)
327 <220> FEATURE:
328 <221> NAME/KEY: BINDING
329 <222> LOCATION: (3)..(3)
330 <223> OTHER INFORMATION: Dpr is bound to a tert-butoxycarbonyl group
332 <220> FEATURE:
333 <221> NAME/KEY: MOD_RES
334 <222> LOCATION: (3)..(3)
335 <223> OTHER INFORMATION: METHYLATION
337 <400> SEQUENCE: 10
W--> 339 Trp Phe Xaa Leu
340 1

RAW SEQUENCE LISTING ERROR SUMMARY DATE: 07/03/2006
PATENT APPLICATION: US/10/537,731 TIME: 11:56:40

Input Set : E:\3874 PTUS sequence listing.txt
Output Set: N:\CRF4\07032006\J537731.raw

Please Note:

Use of n and/or Xaa have been detected in the Sequence Listing. Please review the Sequence Listing to ensure that a corresponding explanation is presented in the <220> to <223> fields of each sequence which presents at least one n or Xaa.

Seq#:1; Xaa Pos. 4 ✓
Seq#:2; Xaa Pos. 4 ✓
Seq#:3; Xaa Pos. 4 ✓
Seq#:4; Xaa Pos. 5 ✓
Seq#:5; Xaa Pos. 5 ✓
Seq#:6; Xaa Pos. 5 ✓
Seq#:7; Xaa Pos. 5 ✓
Seq#:8; Xaa Pos. 5 ✓
Seq#:9; Xaa Pos. 3 ✓
Seq#:10; Xaa Pos. 3 ✓
Seq#:11; Xaa Pos. 4 ✓

VERIFICATION SUMMARY

DATE: 07/03/2006

PATENT APPLICATION: US/10/537,731

TIME: 11:56:40

Input Set : E:\3874 PTUS sequence listing.txt

Output Set: N:\CRF4\07032006\J537731.raw

L:9 M:270 C: Current Application Number differs, Replaced Current Application Number
L:10 M:271 C: Current Filing Date differs, Replaced Current Filing Date
L:42 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:1 after pos.:0
L:77 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:2 after pos.:0
L:107 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 after pos.:0
L:138 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:4 after pos.:0
L:173 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:5 after pos.:0
L:203 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:6 after pos.:0
L:239 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:7 after pos.:0
L:274 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:8 after pos.:0
L:309 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:9 after pos.:0
L:339 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:10 after pos.:0
L:375 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:11 after pos.:0